# This Page Is Inserted by IFW Operations and is not a part of the Official Record

## **BEST AVAILABLE IMAGES**

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images may include (but are not limited to):

- BLACK BORDERS
- TEXT CUT OFF AT TOP, BOTTOM OR SIDES
- FADED TEXT
- ILLEGIBLE TEXT
- SKEWED/SLANTED IMAGES
- COLORED PHOTOS
- BLACK OR VERY BLACK AND WHITE DARK PHOTOS
- GRAY SCALE DOCUMENTS

## IMAGES ARE BEST AVAILABLE COPY.

As rescanning documents will not correct images, please do not report the images to the Image Problem Mailbox.

TRANSMITTAL FORM	Approved for use through 08/30/2003. OMB 0651-0031 U.S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE persons are required to respond to a collection of information unless it displays a valid OMB control number.  Application Number 10/694,075  Filling Date October 27, 2003  First Named Inventor Kandimalla et al.
(to be used for all correspondence after initial filing	
Total Number of Pages in This Submission	ENCLOSURES (Check all that apply)
	Drawing(s)  Licensing-related Papers  Petition  Petition to Convert to a Provisional Application Change of Correspondence Address  Terminal Disclaimer  Request for Refund  CD, Number of CD(s)  Appeal Communication to Board of Appeals and Interferences Appeal Communication to TC (Appeal Notice, Brief, Reply Brief)  Proprietary Information  Status Letter  Other Enclosure(s) (please Identify below):  Remarks  Copy of postcard  PTO Form1 449
	JRE OF APPLICANT, ATTORNEY, OR AGENT
or Individual name  Signature	RTIFICATE OF TRANSMISSION/MAILING  ng facsimile transmitted to the USPTO or deposited with the United States Postal Service with ope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450 on

Typed or printed name	Laura Labier	
Signature	Lausa Laluu	Date 2/5/04/

This collection of information is required by 37 CFR 1.5. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to 12 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.



### IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

**Applicant:** 

Kandimalla et al.

Serial No.:

10/694,075

Filed:

October 27, 2003

**Entitled:** 

Modulation of Immunostimulatory Activity of

Immunostimulatory Oligonucleotide Analogs by Positional

**Chemical Changes** 

**Examiner:** 

NA

**Group Art Unit:** 

NA

Attorney Docket No.:

HYB-005US6 (1006/006)

Commissioner for Patents P. O. Box 1450 Alexandria, VA 22313-1450

#### INFORMATION DISCLOSURE STATEMENT

Sir:

Applicants and their attorney are aware of the following publications and information listed on the attached PTO Form 1449, and in accordance with 37 C.F.R. §1.97 hereby submit these publications for the Examiner's consideration.

Applicants state that the current application is a Divisional application claiming priority to U.S. Patent Application Serial No. 09/965,116, filed September 26, 2001. Applicants also state that the references listed on the attached PTO Form 1449 were previously cited in the parent case and therefore copies of the references are not enclosed herewith.

This submission does not represent that a search has been made and does not constitute an admission that the listed documents are material to patentability or that the listed documents are prior art. If it should be determined that any of the listed documents constitute "prior art" under United States law, Applicants reserve the right to present to the Office relevant facts and law regarding the appropriate status of such documents.

This Information Disclosure Statement is being filed before the mailing date of a first Office Action on the merits and is therefore submitted as both timely and proper. Therefore, no fees are believed to be due.

Respectfully submitted,

Date: 2/5/04

Wayne A. Keown Reg. No. 33,923

Keown & Associates 500 West Cummings Park Suite 1200 Woburn, MA 01801 781-938-1805 (Telephone) 781-938-4777 (Facsimile)

	ł.	5,149,798	09/22/92		Agrawal et al.	536	27		
	INITIAL	DOCUMENT NUMBER	DA	TE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE	
-0171mg									
PANTA	<u> </u>								
TRADE!					1	/27/03		NA	
<u> </u>	Sheet	eet 1 OF 2					,	Group Art Unit	
FEB 0 9	. (بر 2004				<u> </u>				
	ୟ\ <i>(Us</i>	e several shee	ts if nec	essarv)		Kandimalla et al.			
/	5C.	WAY WELL		OIV		Applicant			
INFORMATION DISCLOSURE IN AN APPLICATION					НУ	HYB-005US6 10/694,075			
·	Subt. Form PT	-				Docket Number	App	dication Number	

Foreign Patent Documents							
EXAMINER DOCUMENT DATE COUNTRY CLASS COUNTRY				SUBCLASS	TRANSLATION		
INITIAL	NUMBER				3080000	YES	NO
	WO99/62923		PCT				
		•					
	<u> </u>		L		<u> </u>		

	Other Documents (Including Author, Title, Date Pertinent Pages, Etc.)
C	.   Khorana et al. (1972) "Studies on Polynucleotides," J. Molec, Biol. 72:209
CZ	Reese (1978) "The Chemical Synthesis of Oligo- and Poly-Nucleotides By The Phosporotriester Approach,"  Tetrahedron 34:3143-3179
C	Deoxypolynucleotide Synthesis," Tetrahedron Lett. 22:1859-1862
C4	Connolly et al. (1984) "Synthesis and Characterization of an Octanucleotide Containing the EcoRl Recognition Sequence With A Phosphorothicate Group At The Cleavage Site," Blochemistry 23:3443
C5	Agrawal et al. (1987) "Oligodeoxynucleotise Methylphosphonates: Synthesis and Enzymic Degradation,"  Tetrahedron Lett. 28(31):3539-3542
CE	Blochemistry 27:7237
C7	Immunodeficiency Virus, Proc. Natl. Acad. Sci. USA 85:7079-7083
C8	87-108
Cs	<u>  83:1128-1131                                  </u>
C1	CRC Press, Boca Raton, Florida
<u>C1</u>	
C1	Pisetsky et al. (1994)"Stimulation of Murine Lymphocyte Proliferation By A Phosphorothioate Oligonucleotide With Antisense Activity For Herpes Simplex Virus," 54 Life Sci. 101
C1	Yamamoto et al. (1994) "Lipofection of Synthetic Oligodeoxyribonucleotide Having a Palindromic Sequence of AACGTT to Murine Spenocytes Enhances Interferon Production and Natural Killer Activity," 38 <i>Microbiol. Immunol.</i> 831
C1	Agrawal et al. (1995) "Modified Oligonucleotides as Therapeutic and Diagnostic Agents," Curr. Opin. Biotechnol. 6:12-19
C1	
C1	Vilomon et al. (4000) 90-0 Medite Descentis Destarted DAMA Destarted at the destarted at th
C1	Liang et al. (1996) "Activation of Human B Cells By Phosphorothioate Oligodeoxynucleotides," J. Clin. Invest. 98:1119-1129
C1	<u> </u>
C1	Chu et al. (1997) "CpG Oligodeoxynucleotides Act As Adjuvants That Switch On T Helper 1 (Th1) Immunity," 186 J. Exp. Med. 1623

EXAMINER	DATE CONSIDERED
EXAMINER: Initial if citation is considered, whether or not citati	on is in conformance with MPEP § 609: Draw Line through citation

If not conformance and not considered, include copy with next communication to applicant.

10	Bugt. Form PT	O-1449			Docket Number	Application Number	
<i>(</i> ·	1 5%	PRMATION	DISCLO	SURE	HYB-005US6	10/694,075	
FEB	1 1 12				Applicant		
E	I ÁY	e several she	ets if nece	essary)	Kandimalla e	et al.	
& TRAI	ricy T			<del></del>	Filing Date	Group Art Unit	
<b>~</b>	Sheet	2	OF	2	10/27/03	NA	

·	C20.	Dunford et al. (1997) "Antisense 97: Targeting the Molecular Basis of Disease" (Nature Biotechnology) Conference Abstract, pp. 40
	C21.	Sparwasser et al. (1997) "Macrophages Sense Pathogens Via DNA Motifs: Induction of Turnor Necrosis Factor-α-Mediated Shock," 27 Eur. J. Immunol. 1671
	C22.	Zhao et al. (1997) "Pattern and Kinetics of Cytokine Production Following Administration of Phosphorothioate Oligonucleotides in Mice," 7 Antisense Nucleic Acid Drug. Dev. 495
	C23.	McCluside et al. (1998) "Cutting Edge: CpG DNA is A Potent Enhancer of Systemic and Mucosal immune Responses Against Hepatitis B Surface Antigen with Intranasal Administration to Mice," J. Immunol. 161:4463-4466
	C24.	Moldoveanu et al. (1998) "CpG DNA, A Novel Immune Enhancer for Systemic and Mucosal Immunization With Influenza Virus." Vaccine 16:1216-1224
	C25.	Sparwasser et al. (1998) "Bacterial DNA and Immunostimulatory CpG Oligonucleotides Trigger Maturation and ACtivation of Murine Dendritic Cells," 28 Eur. J. Immunol. 2045
	C26.	Tokunaga et al. (1999) "How BCG Led to the Discovery of Immunostimulatory DNA," 52 Jap. J. Infect. Dis. 1
	C27.	Zhao et al. (1999) "Site of Chemical Modifications in CpG Containing Phosphorothicate Oligodecxynucleotide Modulates its Immunostimulatory Activity," Bloorg. & Med. Chem. Lett. 9:3453-3458
	C28.	Agrawal et al. (2000) "Antisense Therapeutics: Is It As Simple As Complementary Base Recognition," 6 Mol. Med. Today 72
	C29.	Zhao et al. (2000) "Immunostimulatory Activity of CpG Containing Phosphorothioate Oligodeoxynucleotide is Modulated by Modification of a Single Deoxynucleoside," <i>Bloorg. &amp; Med. Chem. Lett.</i> 10:1051-1054
	LC30	Agrawal et al., "Antisense therapeutics", Curr. Opin.Chem.
		Biol., 2:519-528, 1998.
	C31	Chaix et al., "3'-3' Linked Oligonucleotides: Synthesis and
	1.23	Charlet at 1 2 5 5 1 1 1 2 1 1 1 1 2 1 1 1 1 1 1 1
	1 224	Stability Studies", Biorg. & Med.Chem., 6:827-832, 1996.
	C32	
		Godeoxynucleotides", Antisense & Nucl. Acid Drug Dev
		8:181-184, 1998.
	C33	Yu et al., "Accessible 5'-End of CpG-Containing",
	C34	
	<del>  ~~  </del>	Riographia et al., "Effect of Chemical Modifications ",
		Bioorganic & Medicinal Chemistry, 9:807-813, 2001.
	C35	International Search Report (PCT APP. No. PCT/USO1/30137)
<del></del>	<del></del>	
<b></b>	<del>                                     </del>	
	ļ	
	لــــــا	
	L	
	<del>  </del>	
<b> </b>	<del>  </del>	
	<del>  </del>	
	——	
L	1	

EXAMINER	DATE CONSIDERED
TVANIAITD	

EXAMINER: Initial if citation is considered, whether or not citation is in conformance with MPEP § 609: Draw Line through citation if not conformance and not considered. Include copy with next communication to applicant.